SEQUENCE LISTING

```
<110> Duprat, Fabrice
        Lesage, Florian
        Fink, Michel
        Lazdunski, Michel
  <120> FAMILY OF MAMMALIAN POTASSIUM CHANNELS, THEIR CLONING
        AND THEIR USE, ESPECIALLY FOR THE SCREENING OF DRUGS
  <130> 1201-CIP-DIV-00
  <141> 2001-08-24
  <150> 09/144,914
  <151> 1998-09-01
  <150> 08/749,816
  <151> 1996-11-15
  <150> 60/095,234
  <151> 1998-08-04
  <150> FR 96/01565
a <151> 1996-02-08
ıI)
  <160> 24
Ð

↓ <170> PatentIn Ver. 2.0

Ð
<u>_</u> <210> 1
(211> 1894)
  <212> DNA
  <213> Homo sapiens
<220>
(1) <221> CDS
<222> (183)..(1190)
<220>
☐ <223> TWIK-1
<400> 1
  gegggegga geeaggeeg ggeggggeg gggeggegg ggeeagaaga ggeggeggge 120
  cgcgctccgg ccggtctgcg gcgttggcct tggctttggc tttggcggcg gcggtggaga 180
                                                                  227
  ag atg ctg cag tcc ctg gcc ggc agc tcg tgc gtg cgc ctg gtg gag
     Met Leu Gln Ser Leu Ala Gly Ser Ser Cys Val Arg Leu Val Glu
       1
  cgg cac cgc tcg gcc tgg tgc ttc ggc ttc ctg gtg ctg ggc tac ttg
  Arg His Arg Ser Ala Trp Cys Phe Gly Phe Leu Val Leu Gly Tyr Leu
                  20
  ctc tac ctg gtc ttc ggc gca gtg gtc ttc tcc tcg gtg gag ctg ccc
                                                                  323
  Leu Tyr Leu Val Phe Gly Ala Val Val Phe Ser Ser Val Glu Leu Pro
               35
```

								gag Glu 55									371
								tct Ser									419
								aac Asn									467
								gac Asp									515
								ggt Gly									563
								atc Ile 135									611
	Phe							gct Ala									659
	Val 160	acc Thr	cgc Arg	agg Arg	ccg Pro	gtc Val 165	ctc Leu	tac Tyr	ttc Phe	cac His	atc Ile 170	cgc Arg	tgg Trp	ggc Gly	ttc Phe	tcc Ser 175	707
	aag Lys							cat His									755
	gtg Val	tcc Ser	tgc Cys	ttc Phe 195	ttc Phe	ttc Phe	atc Ile	ccg Pro	gcc Ala 200	gct Ala	gtc Val	ttc Phe	tca Ser	gtc Val 205	ctg Leu	gag Glu	803
								tcc Ser 215									851
								tat Tyr									899
								att Ile									947
								gtt Val									995
								atg Met									1043
	gag	gat	cag	gtg	cac	atc	ata	gag	cat	gac	caa	ctg	tcc	ttc	tcc	tcg	1091

```
290
                             295
  atc aca gac cag gca gct ggc atg aaa gag gac cag aag caa aat gag
                                                                   1139
  Ile Thr Asp Gln Ala Ala Gly Met Lys Glu Asp Gln Lys Gln Asn Glu
      305
  cet ttt qtq qee ace caq tea tet qee tge gtq gat gge cet gea aac
                                                                   1187
  Pro Phe Val Ala Thr Gln Ser Ser Ala Cys Val Asp Gly Pro Ala Asn
  320
                     325
                                         330
                                                                   1240
  cat tgagcgtagg atttgttgca ttatgctaga gcaccagggt cagggtgcaa
  ggaagaggct taagtatgtt catttttatc agaatgcaaa agcgaaaatt atgtcacttt 1300
  aagaaatagc tactgtttgc aatgtcttat taaaaaacaa caaaaaaaga cacatggaac 1360
  aaaqaaqctg tgaccccagc aggatgtcta atatgtgagg aaatgagatg tccacctaaa 1420
  attcatatgt gacaaaatta tctcgacctt acataggagg agaatacttg aagcagtatg 1480
ctqctqtqqt taqaaqcaqa ttttatactt ttaactggaa actttggggt ttgcatttag 1540
🔁 atcatttagc tgatggctaa atagcaaaat ttatatttag aagcaaaaaa aaaaagcata 1600
Ü
  gagatgtgtt ttataaatag gtttatgtgt actggtttgc atgtacccac ccaaaatgat 1660
  tatttttgga gaatctaagt caaactcact atttataatg cataggtaac cattaactat 1720
🗸 gtacatataa agtataaata tgtttatatt ctgtacatat ggtttaggtc accagatcct 1780
agtgtagttc tgaaactaag actatagata ttttgtttct tttgatttct ctttatacta 1840
  [] <210> 2
<u>=</u> <211> 336
  <212>.PRT
  <213> Homo sapiens
  <220>
  <223> TWIK-1
  <400> 2
  Met Leu Gln Ser Leu Ala Gly Ser Ser Cys Val Arg Leu Val Glu Arg
  His Arg Ser Ala Trp Cys Phe Gly Phe Leu Val Leu Gly Tyr Leu Leu
  Tyr Leu Val Phe Gly Ala Val Val Phe Ser Ser Val Glu Leu Pro Tyr
           35
  Glu Asp Leu Leu Arg Gln Glu Leu Arg Lys Leu Lys Arg Arg Phe Leu
  Glu Glu His Glu Cys Leu Ser Glu Gln Gln Leu Glu Gln Phe Leu Gly
```

Glu Asp Gln Val His Ile Ile Glu His Asp Gln Leu Ser Phe Ser Ser

Arg Val Leu Glu Ala Ser Asn Tyr Gly Val Ser Val Leu Ser Asn Ala Ser Gly Asn Trp Asn Trp Asp Phe Thr Ser Ala Leu Phe Phe Ala Ser 105 Thr Val Leu Ser Thr Thr Gly Tyr Gly His Thr Val Pro Leu Ser Asp Gly Gly Lys Ala Phe Cys Ile Ile Tyr Ser Val Ile Gly Ile Pro Phe Thr Leu Leu Phe Leu Thr Ala Val Val Gln Arg Ile Thr Val His Val Thr Arg Arg Pro Val Leu Tyr Phe His Ile Arg Trp Gly Phe Ser Lys Gln Val Val Ala Ile Val His Ala Val Leu Leu Gly Phe Val Thr Val Ser Cys Phe Phe Phe Ile Pro Ala Ala Val Phe Ser Val Leu Glu Asp 🖺 Asp Trp Asn Phe Leu Glu Ser Phe Tyr Phe Cys Phe Ile Ser Leu Ser Û 210 Q, Thr Ile Gly Leu Gly Asp Tyr Val Pro Gly Glu Gly Tyr Asn Gln Lys Ü FPhe Arg Glu Leu Tyr Lys Ile Gly Ile Thr Cys Tyr Leu Leu Gly D Leu Ile Ala Met Leu Val Val Leu Glu Thr Phe Cys Glu Leu His Glu Leu Lys Lys Phe Arg Lys Met Phe Tyr Val Lys Lys Asp Lys Asp Glu 275 Asp Gln Val His Ile Ile Glu His Asp Gln Leu Ser Phe Ser Ser Ile 295 Thr Asp Gln Ala Ala Gly Met Lys Glu Asp Gln Lys Gln Asn Glu Pro Phe Val Ala Thr Gln Ser Ser Ala Cys Val Asp Gly Pro Ala Asn His

```
<210> 3
```

<211> 2514

<212> DNA

<213> Homo sapiens

<220>₄

<221> CDS

<222> (126)..(1307)

<220>

<223> TASK

<400> 3 tgccctgcgc ggatagcggc gagcgcagcc atgccccagg ccgcctccgg ggcagcagca 60 gcggcggccg gggccgatgc gcgggccggg ggccggggg ggccggcggc ggcccgggcg 120 ggacg atg aag cgg cag aac gtg cgc acg ctg gcg ctc atc gtg tgc acc 170 Met Lys Arq Gln Asn Val Arq Thr Leu Ala Leu Ile Val Cys Thr ttc acc tac ctg ctg gtg ggc gcc gcg gtc ttc gac gcg ctg gag tcg 218 Phe Thr Tyr Leu Leu Val Gly Ala Ala Val Phe Asp Ala Leu Glu Ser gag ccc gag ctg atc gag cgg cag cgg ctg gag ctg cgg cag cag gag Glu Pro Glu Leu Ile Glu Arg Gln Arg Leu Glu Leu Arg Gln Glu 35 ctg cqq qcq cqc tac aac ctc agc cag ggc ggc tac gag gag ctg gag Leu Arg Ala Arg Tyr Asn Leu Ser Gln Gly Gly Tyr Glu Glu Leu Glu 50 cgc gtc gtg ctg cgc ctc aag ccg cac aag gcc ggc gtg cag tgg cgc Arg Val Val Leu Arg Leu Lys Pro His Lys Ala Gly Val Gln Trp Arg 65 intto goo ggo too tto tao tto goo ato aco gto ato aco aco ato ggo 410 Phe Ala Gly Ser Phe Tyr Phe Ala Ile Thr Val Ile Thr Thr Ile Gly 85 80 W 🚇 tac ggg cac gcg gca ccc agc acg gat ggc ggc aag gtg ttc tgc atg 458 正Tyr Gly His Ala Ala Pro Ser Thr Asp Gly Gly Lys Val Phe Cys Met 100 110 Ħ ttc tac gcg ctg ctg ggc atc ccg ctc acg ctc gtc atg ttc cag agc 506 Phe Tyr Ala Leu Leu Gly Ile Pro Leu Thr Leu Val Met Phe Gln Ser 125 115 Mictg ggc gag cgc atc aac acc ttg gtg agg tac ctg ctg cac cgc gcc 554 Leu Gly Glu Arg Ile Asn Thr Leu Val Arg Tyr Leu Leu His Arg Ala 130 135 140 📇 aag aag ggg ctg ggc atg cgg cgc gcc gac gtg tcc atg gcc aac atg 602 Lys Lys Gly Leu Gly Met Arg Arg Ala Asp Val Ser Met Ala Asn Met 145 150 650 gtg ctc atc ggc ttc ttc tcg tgc atc agc acg ctg tgc atc ggc gcc Val Leu Ile Gly Phe Phe Ser Cys Ile Ser Thr Leu Cys Ile Gly Ala 175 160 165 698 gcc gcc ttc tcc cac tac gag cac tgg acc ttc ttc cag gcc tac tac Ala Ala Phe Ser His Tyr Glu His Trp Thr Phe Phe Gln Ala Tyr Tyr 180 190 185 tac tgc ttc atc acc ctc acc acc atc ggc ttc ggc gac tac gtg gcg 746 Tyr Cys Phe Ile Thr Leu Thr Thr Ile Gly Phe Gly Asp Tyr Val Ala 195 205 ctg cag aag gac cag gcc ctg cag acg cag ccg cag tac gtg gcc ttc Leu Gln Lys Asp Gln Ala Leu Gln Thr Gln Pro Gln Tyr Val Ala Phe 210 215 220

	agc Ser	ttc Phe 225	gtc Val	tac Tyr	atc Ile	ctt Leu	acg Thr 230	ggc Gly	ctc Leu	acg Thr	gtc Val	atc Ile 235	ggc Gly	gcc Ala	ttc Phe	ctc Leu	842
	aac Asn 240	ctc Leu	gtg Val	gtg Val	ctg Leu	cgc Arg 245	ttc Phe	atg Met	acc Thr	atg Met	aac Asn 250	gcc Ala	gag Glu	gac Asp	gag Glu	aag Lys 255	890
		gac Asp															938
	ggc Gly	ggc Gly	gga Gly	ggg Gly 275	ggt Gly	ggc Gly	agc Ser	gcg Ala	cac His 280	act Thr	acg Thr	gac Asp	acc Thr	gcc Ala 285	tca Ser	tcc Ser	986
	acg Thr	gcg Ala	gca Ala 290	gcg Ala	ggc Gly	ggc Gly	ggc Gly	ggc Gly 295	ttc Phe	cgc Arg	aac Asn	gtc Val	tac Tyr 300	gcg Ala	gag Glu	gtg Val	1034
		cac His 305															1082
	aag Lys 320	ctg Leu	cag Gln	tac Tyr	tcc Ser	atc Ile 325	ccc Pro	atg Met	atc Ile	atc Ile	ccg Pro 330	cgg Arg	gac Asp	ctc Leu	tcc Ser	acg Thr 335	1130
		gac Asp															1178
	Arg	tac Tyr	Ser	Asp 355	Thr	Pro	Ser	Arg	Arg 360	Cys	Leu	Cys	Ser	Gly 365	Ala	Pro	1226
	Arg	tcc Ser	370	ile	Ser	Ser	vai	375	Thr	GTÀ	Leu	HIS	380	Leu	ser	Thr	1274
	ttc Phe	cgc Arg 385	ggc Gly	ctc Leu	atg Met	aag Lys	cgc Arg 390	agg Arg	agc Ser	tcc Ser	gtg Val	tgad	etgeo	cc d	gaggg	gacctg	1327
	gago	cacct	gg g	gggcg	cggç	ac do	ggga	accco	tgo	tgctgggagg			gaga	1387			
1	cctt	ctgo	cc a	agtgo	gaco	cc cg	caca	aacat	ccc	ctcac	ccac	tctc	cccc	cag o	cacco	ccatc	1447
	tcc	gacto	gtg d	cctgcttgca ccagccggca					. ggaggccggg			ctctgaggac ccctggggcc				1507	
	ccca	atcgg	gag d	ccctgcaaat tccg				gaaat gtgaaacttg			gtggggtcag ggaggaaagg					1567	
	caga	agct	gg g	gagco	ctccc	ct to	cctt	ctttgaa		aatctaagaa		gctcccagtc ctca			ctcag	gagacc	1627
	ctgo	ctggt	ac c	cacac	ccca	ac ct	tcgg	gagg	ggad	gacttcatgt			gtgta	1687			
	tatt	tata	acc t	tctgtcctgc ta				ctccc	aco	accttccctt			ggttccaaaa gccagggtgt			ggtgt	1747
	ctat	gtco	caa g	aa gtcaccccta			cago	ccca	a cto	ctcccttcc			tcatccccag ctgtgtctc			gtctcc	1807
	caac	cctcc	ct t	cgto	ıttgt	t tt	gcat	ggct	ttç	gcagt	tat	ggag	gaaaç	gtg g	gaaac	ccagc	1867

.

:

-

agtecetaaa getggteece agaaageagg acagaaagaa ggagggacag geaggeagea 1927
ggaggggega getgggagge aggaggeage ggeetgteag tetgcagaat ggtegeactg 1987
gaggtteaag etaactggee tecageeaca tteteatage aggtaggaet teageettee 2047
agacactgee ettagaatet ggaacagaag actteagaet caccataatt getgataatt 2107
acceactett aaatttgteg agtgatttt ageetetgaa aactetatge tggeeactga 2167
tteetttgag teteacaaaa ecetaettag gteateaggg eaggagttet cacteceatt 2227
ttacagatga acetgtatte ecaacacttt tggaggetga ggttggagga ttgettgage 2287
ceaggaatte gagaceagee taggtgacat agtgagacee eatetetaca aaaaataaaa 2347
aattaaceag gtgtggtgge aegtgeetgg gagteecage gaettgggag getgaggtgg 2407
gaggattgtt tgageetggg aggtegagge tgtagtgage eetgattgea ecactgtaet 2467
ceageetggg tgacagggea agaceetgte teaaaaaaaa aaaaaaa

3<210> 4 一<211> 394 直<212> PRT <213> Homo sapiens **₽**<220> . <223> TASK **4**<400> 4 Met Lys Arg Gln Asn Val Arg Thr Leu Ala Leu Ile Val Cys Thr Phe 🌣 Thr Tyr Leu Leu Val Gly Ala Ala Val Phe Asp Ala Leu Glu Ser Glu 20 Pro Glu Leu Ile Glu Arg Gln Arg Leu Glu Leu Arg Gln Glu Leu Arg Ala Arg Tyr Asn Leu Ser Gln Gly Gly Tyr Glu Glu Leu Glu Arg Val Val Leu Arg Leu Lys Pro His Lys Ala Gly Val Gln Trp Arg Phe Ala Gly Ser Phe Tyr Phe Ala Ile Thr Val Ile Thr Thr Ile Gly Tyr Gly His Ala Ala Pro Ser Thr Asp Gly Gly Lys Val Phe Cys Met Phe Tyr Ala Leu Leu Gly Ile Pro Leu Thr Leu Val Met Phe Gln Ser Leu Gly Glu Arg Ile Asn Thr Leu Val Arg Tyr Leu Leu His Arg Ala Lys Lys Gly Leu Gly Met Arg Arg Ala Asp Val Ser Met Ala Asn Met Val

Leu Ile Gly Phe Phe Ser Cys Ile Ser Thr Leu Cys Ile Gly Ala Ala 165 170 175

Ala Phe Ser His Tyr Glu His Trp Thr Phe Phe Gln Ala Tyr Tyr 180 185 190

Cys Phe Ile Thr Leu Thr Thr Ile Gly Phe Gly Asp Tyr Val Ala Leu 195 200 205

Gln Lys Asp Gln Ala Leu Gln Thr Gln Pro Gln Tyr Val Ala Phe Ser 210 215 220

Phe Val Tyr Ile Leu Thr Gly Leu Thr Val Ile Gly Ala Phe Leu Asn 225 235 240

Leu Val Val Leu Arg Phe Met Thr Met Asn Ala Glu Asp Glu Lys Arg
245 250 255

Asp Ala Glu His Arg Ala Leu Leu Thr Arg Asn Gly Gln Ala Gly Gly 260 265 270

Gly Gly Gly Gly Ser Ala His Thr Thr Asp Thr Ala Ser Ser Thr 275 280 285

Ala Ala Ala Gly Gly Gly Phe Arg Asn Val Tyr Ala Glu Val Leu 295 300

Leu Gln Tyr Ser Ile Pro Met Ile Ile Pro Arg Asp Leu Ser Thr Ser 325 330 335

Asp Thr Cys Val Glu Gln Ser His Ser Ser Pro Gly Gly Gly Arg
340 345 350

Tyr Ser Asp Thr Pro Ser Arg Arg Cys Leu Cys Ser Gly Ala Pro Arg
355
360
365

Ser Ala Ile Ser Ser Val Ser Thr Gly Leu His Ser Leu Ser Thr Phe 370 375 380

Arg Gly Leu Met Lys Arg Arg Ser Ser Val 385 390

<210> 5

145

<211> 405

<212> PRT

<213> Murine

<220>

<223> TASK

<400> 5 ·

Glu Asn Val Arg Thr Leu Ala Leu Ile Val Cys Thr Phe Thr Tyr Leu 1 5 10 15

Leu Val Gly Ala Ala Val Phe Asp Ala Leu Glu Ser Glu Pro Glu Met

Ile Glu Arg Gln Arg Leu Glu Leu Arg Gln Leu Glu Leu Arg Ala Arg Tyr Asn Leu Ser Glu Gly Gly Tyr Glu Glu Leu Glu Arg Val Val Leu Arg Leu Lys Pro His Lys Ala Gly Val Gln Trp Arg Phe Ala Gly Ser Phe Tyr Phe Ala Ile Thr Val Ile Thr Thr Ile Gly Tyr Gly His Ala Ala Pro Ser Thr Asp Gly Gly Lys Val Phe Cys Met Phe Tyr Ala Leu Leu Gly Ile Pro Leu Thr Leu Ile Met Phe Gln Ser Leu Gly Glu Arg 120 Ile Asn Thr Phe Val Arg Tyr Leu Leu His Arg Ala Lys Arg Gly Leu 135 Gly Met Arg His Ala Glu Val Ser Met Ala Asn Met Val Leu Ile Gly 150 引 Phe Val Ser Cys Ile Ser Thr Leu Cys Ile Gly Ala Ala Ala Phe Ser 165 Ш Tyr Tyr Glu Arg Trp Thr Phe Phe Gln Ala Tyr Tyr Tyr Cys Phe Ile 180 185 Ò #Thr Leu Thr Thr Ile Gly Phe Gly Asp Tyr Val Ala Leu Gln Lys Asp 195 33 Gln Ala Leu Gln Thr Gln Pro Gln Tyr Val Ala Phe Ser Phe Val Tyr 215 M .∰Ile Leu Thr Gly Leu Thr Val Ile Gly Ala Phe Leu Asn Leu Val Val **225** Leu Arg Phe Met Thr Met Asn Ala Glu Asp Glu Lys Arg Asp Ala Glu His Arg Ala Leu Leu Thr His Asn Gly Gln Ala Val Gly Leu Gly Gly Leu Ser Cys Leu Ser Gly Ser Leu Gly Asp Val Arg Pro Arg Asp Pro Val Thr Cys Ala Ala Ala Ala Gly Gly Val Gly Val Gly Gly 295 Ser Gly Phe Arg Asn Val Tyr Ala Glu Val Leu His Phe Gln Ser Met Cys Ser Cys Leu Trp Tyr Lys Ser Arg Glu Lys Leu Gln Tyr Ser Ile Pro Met Ile Ile Pro Arg Asp Leu Ser Thr Ser Asp Thr Cys Val Glu 340 345

His Ser His Ser Ser Pro Gly Gly Gly Gly Arg Tyr Ser Asp Thr Pro Ser His Pro Cys Leu Cys Ser Gly Thr Gln Arg Ser Ala Ile Ser Ser Val Ser Thr Gly Leu His Ser Leu Ala Ala Phe Arg Gly Leu Met Lys Arg Arg Ser Ser Val <210> 6 <211> 347 <212> PRT <213> Unknown <223> Description of Unknown Organism: TWIK-1 homolog Met Tyr Thr Asp Glu Gly Glu Tyr Ser Gly Asp Thr Asp His Gly Gly Ser Thr Met Gln Lys Met Ser Pro Asn Thr Arg Gln Asn Phe Arg Gln Asn Val Asn Val Val Val Cys Leu Ser Ala Ala Ile Thr Leu Leu Val Phe Asn Leu Ile Gly Ala Gly Ile Phe Tyr Leu Ala Glu Thr Gln Asn Ser Ser Glu Ser Leu Asn Glu Asn Ser Glu Val Ser Lys Cys Leu His Asn Leu Pro Ile Gly Gly Lys Ile Thr Ala Glu Met Lys Ser Lys Leu 85 Gly Lys Cys Leu Thr Lys Ser Ser Arg Ile Asp Gly Phe Gly Lys Ala Ile Phe Phe Ser Trp Thr Leu Tyr Ser Thr Val Gly Tyr Gly Ser Leu 115 Tyr Pro His Ser Thr Leu Gly Arg Tyr Leu Thr Ile Phe Tyr Ser Leu Leu Met Ile Pro Val Phe Ile Ala Phe Lys Phe Glu Phe Gly Thr Phe Leu Ala His Phe Leu Val Val Val Ser Asn Arg Thr Arg Leu Ala Val Lys Lys Ala Tyr Tyr Lys Leu Ser Gln Asn Pro Glu Asn Ala Glu Thr 180

Pro Ser Asn Ser Leu Gln His Asp Tyr Leu Ile Phe Leu Ser Ser Leu

205

200

195

```
Leu Leu Cys Ser Ile Ser Leu Leu Ser Ser Ser Ala Leu Phe Ser Ser
  Ile Glu Asn Ile Ser Tyr Leu Ser Ser Val Tyr Phe Gly Ile Ile Thr
  225
  Met Phe Leu Ile Gly Ile Gly Asp Ile Val Pro Thr Asn Leu Val Trp
  Phe Ser Gly Tyr Cys Met Leu Phe Leu Ile Ser Asp Val Leu Ser Asn
  Gln Ile Phe Tyr Phe Cys Gln Ala Arg Val Arg Tyr Phe Phe His Ile
  Leu Ala Arg Lys Ile Leu Leu Arg Glu Glu Asp Asp Gly Phe Gln
  Leu Glu Thr Thr Val Ser Leu Gln His Ile Pro Ile Ile Asn Ser Gln
  Cys Met Pro Ser Leu Val Leu Asp Cys Glu Lys Glu Glu Leu Asp Asn
                 325
Asp Glu Lys Leu Ile Ser Ser Leu Thr Ser Thr
             340
Ū
W
□<210> 7
₹<211> 383
靐<213> Unknown
  <223> Description of Unknown Organism: TWIK-1 homolog
型<400> 7
靐Met Thr Val Ser Met Glu Glu Asn Ser Lys Ile Gln Met Leu Ser Ala
                                                           15
  Thr Ser Lys Asp Lys Lys Val Ala Thr Asp Arg Ser Leu Leu Asn Lys
                                   25
  Tyr His Leu Gly Pro Leu Ala Leu His Thr Gly Leu Val Leu Ser Cys
           35
                                                   45
  Val Thr Tyr Ala Leu Gly Gly Ala Tyr Leu Phe Leu Ser Ile Glu His
  Pro Glu Glu Leu Lys Arg Arg Glu Lys Ala Ile Arg Glu Phe Gln Asp
  Leu Lys Gln Gln Phe Met Gly Asn Ile Thr Ser Gly Ile Glu Asn Ser
  Glu Gln Ser Ile Glu Ile Tyr Thr Lys Lys Leu Ile Leu Met Leu Glu
             100
                                  105
  Asp Ala His Asn Ala His Ala Phe Glu Tyr Phe Phe Leu Asn His Glu
                              120
                                                  125
         115
```

Thr Thr Val Ile Pro Val Gly Tyr Gly Tyr Ile Phe Pro Val Ser Ala Tyr Gly Arg Met Cys Leu Ile Ala Tyr Ala Leu Leu Gly Ile Pro Leu Thr Leu Val Thr Met Ala Asp Thr Gly Lys Phe Ala Ala Gln Leu Val Thr Arg Trp Phe Gly Asp Asn Asn Met Ala Ile Pro Ala Ala Ile Phe Val Cys Leu Leu Phe Ala Tyr Pro Leu Val Val Gly Phe Ile Leu Cys 210 215 Ser Thr Ser Asn Ile Thr Tyr Leu Asp Ser Val Tyr Phe Ser Leu Thr Ser Ile Phe Thr Ile Gly Phe Gly Asp Leu Thr Pro Asp Met Asn Val 245 Ile His Met Val Leu Phe Leu Ala Val Gly Val Ile Leu Val Thr Ile 265 Ф Thr Leu Asp Ile Val Ala Ala Glu Met Ile Asp Arg Val His Tyr Met 275 🛱 Gly Arg His Val Gly Lys Ala Lys Glu Leu Ala Gly Lys Met Phe Gln 295 Leu Ala Gln Ser Leu Asn Met Lys Gln Gly Leu Val Ser Gly Val Gly 320 310 Ō Gln Leu His Ala Leu Ala Arg Phe Gly Met Leu Val Gly Arg Glu Glu 325 Val Asp Lys Thr Gln Glu Asp Gly Ile Ile Ala Phe Ser Pro Asp Val 340 Met Asp Gly Leu Glu Phe Met Asp Thr Leu Ser Ile Tyr Ser Arg Arg 360 Ser Arg Arg Ser Ala Glu Asn Ser Ala Arg Asn Leu Phe Leu Ser 370 375 <210> 8 <211> 370 <212> PRT <213> Murine <220> <223> TREK-1

Met Ala Ala Pro Asp Leu Leu Asp Pro Lys Ser Ala Ala Gln Asn Ser

10

Ile Pro Lys Asp Met Trp Thr Phe Ser Ser Ala Leu Val Phe Thr Thr

Lys Pro Arg Leu Ser Phe Ser Ser Lys Pro Thr Val Leu Ala Ser Arg Val Glu Ser Asp Ser Ala Ile Asn Val Met Lys Trp Lys Thr Val Ser Thr Ile Phe Leu Val Val Val Leu Tyr Leu Ile Ile Gly Ala Ala Val Phe Lys Ala Leu Glu Gln Pro Gln Glu Ile Ser Gln Arg Thr Thr Ile Val Ile Gln Lys Gln Thr Phe Ile Ala Gln His Ala Cys Val Asn Ser Thr Glu Leu Asp Glu Leu Ile Gln Gln Ile Val Ala Ala Ile Asn Ala Gly Ile Ile Pro Leu Gly Asn Ser Ser Asn Gln Val Ser His Trp Asp Leu Gly Ser Ser Phe Phe Phe Ala Gly Thr Val Ile Thr Thr Ile Gly phe Gly Asn Ile Ser Pro Arg Thr Glu Gly Gly Lys Ile Phe Cys Ile Ile Tyr Ala Leu Leu Gly Ile Pro Leu Glu Gly Phe Leu Leu Ala Gly 165 🔛 Val Gly Asp Gln Leu Gly Thr Ile Phe Gly Lys Gly Ile Ala Lys Val 185 Glu Asp Thr Phe Ile Lys Trp Asn Val Ser Gln Thr Lys Ile Arg Ile 195 🏮 Ile Ser Thr Ile Ile Phe Ile Leu Phe Gly Cys Val Leu Phe Val Ala Leu Pro Ala Val Ile Phe Lys His Ile Glu Gly Trp Ser Ala Leu Asp Ala Ile Tyr Phe Val Val Ile Thr Leu Thr Thr Ile Gly Phe Gly Asp 250 Tyr Val Ala Gly Gly Ser Asp Ile Glu Tyr Leu Asp Phe Tyr Lys Pro Val Val Trp Phe Trp Ile Leu Val Gly Leu Ala Tyr Phe Ala Ala Val Leu Ser Met Ile Gly Asp Trp Leu Arg Val Ile Ser Lys Lys Thr Lys Glu Glu Val Gly Glu Phe Arg Ala His Ala Ala Glu Trp Thr Ala Asn Val Thr Ala Glu Phe Lys Glu Thr Arg Arg Arg Leu Ser Val Glu Ile

```
Tyr Asp Lys Phe Gln Arg Ala Thr Ser Val Lys Arg Lys Leu Ser Ala
                                  345
  Glu Leu Ala Gly Asn His Asn Gln Glu Leu Thr Pro Cys Met Arg Thr
                              360
  Cys Leu
      370
  <210> 9
  <211> 27
  <212> PRT
  <213> Homo sapiens
  <220>
  <223> TWIK-1 P1
  <400> 9
  Phe Thr Ser Ala Leu Phe Phe Ala Ser Thr Val Leu Ser Thr Thr Gly
  Tyr Gly His Thr Val Pro Leu Ser Asp Gly Gly
            . 20
ıΦ
₫<210> 10
(211> 27
___<212> PRT
风<220>
.⊈<223> TWIK-1 P2
400> 10
Phe Leu Glu Ser Phe Tyr Phe Cys Phe Ile Ser Leu Ser Thr Ile Gly
Leu Gly Asp Tyr Val Pro Gly Glu Gly Tyr Asn
              20
                                  25
|=
 <210> 11
 <211> 27
  <212> PRT
 <213> Unknown
 <220>
  <223> Description of Unknown Organism: P domain of
       representative K+ channel sequence
  <220>
 <223> TOK-1 P2
 <400> 11
 Tyr Phe Asn Cys Ile Tyr Phe Cys Phe Leu Cys Leu Leu Thr Ile Gly
 Tyr Gly Asp Tyr Ala Pro Arg Thr Gly Ala Gly
              20
```

```
<210> 12
  <211> 27
  <212> PRT
· <213> Unknown
  <220>
  <223> Description of Unknown Organism: P domain of
        representative K+ channel sequence
  <220>
  <223> TOK-1 P1
  <400> 12
  Tyr Gly Asn Ala Leu Tyr Phe Cys Thr Val Ser Leu Leu Thr Val Gly
  Leu Gly Asp Ile Leu Pro Lys Ser Val Gly Ala
               20
  <210> 13
  <211> 27
  <212> PRT
=<213> Unknown
Ē
4<220>
223> Description of Unknown Organism: P domain of
        representative K+ channel sequence
Ü
(223> Slo
# <400> 13
Tyr Trp Thr Cys Val Tyr Phe Leu Ile Val Thr Met Ser Thr Val Gly
Ď
Tyr Gly Asp Val Tyr Cys Glu Thr Val Leu Gly
S
               20
  <210> 14
  <211> 27
  <212> PRT
  <213> Unknown
  <223> Description of Unknown Organism: P domain of
       representative K+ channel sequence
  <220>
  <223> Shaker
  <400> 14
  Ile Pro Asp Ala Phe Trp Trp Ala Val Val Thr Met Thr Thr Val Gly
   1
                                       10
  Tyr Gly Asp Met Thr Pro Val Gly Phe Trp Gly
               20
```

```
<210> 15
  <211> 27
  <212> PRT
  <213> Unknown
  <220>
  <223> Description of Unknown Organism: P domain of
        representative K+ channel sequence
  <220>
  <223> Shab
  <400> 15
  Ile Pro Glu Ala Phe Trp Trp Ala Gly Ile Thr Met Thr Thr Val Gly
  Tyr Gly Asp Ile Cys Pro Thr Thr Ala Leu Gly
               20
  <210> 16
  <211> 27
  <212> PRT
  <213> Unknown
:□<220>
ú <223> Description of Unknown Organism: P domain of
        representative K+ channel sequence
₾<220>
-<223> Shal
Ď
<u></u> <400> 16
" Ile Pro Ala Ala Phe Trp Tyr Thr Ile Val Thr Met Thr Thr Leu Gly
Tyr Gly Asp Met Val Pro Glu Thr Ile Ala Gly
               20
-
  <210> 17
  <211> 27
  <212> PRT
  <213> Unknown
  <220>
  <223> Description of Unknown Organism: P domain of
        representative K+ channel sequence
  <220>
  <223> Shaw
  <400> 17
  Ile Pro Leu Gly Leu Trp Trp Ala Leu Val Thr Met Thr Thr Val Gly
  Tyr Gly Asp Met Ala Pro Lys. Thr Tyr Ile Gly
                                    25
               20
```

```
<211> 27
  <212> PRT
  <213> Unknown
  <220>
  <223> Description of Unknown Organism: P domain of
        representative K+ channel sequence
  <220>
  <223> KAT1
  <400> 18
  Tyr Val Thr Ala Leu Tyr Trp Ser Ile Thr Thr Leu Thr Thr Gly
  Tyr Gly Asp Phe His Ala Glu Asn Pro Arg Glu
  <210> 19
  <211> 27
  <212> PRT
  <213> Unknown
U<220>
₫<223> Description of Unknown Organism: P domain of
        representative K+ channel sequence
Ф
W
(220)
  <223> AKT1
<sup>™</sup><400> 19
#Tyr Val Thr Ser Met Tyr Trp Ser Ile Thr Thr Leu Thr Thr Val Gly
   1
                                                             15
Į;
Tyr Gly Asp Ile His Pro Val Asn Thr Lys Glu
               20
                                    25
17.1
<u> |</u><210> 20
<211> 27
  <212> PRT
  <213> Unknown
  <223> Description of Unknown Organism: P domain of
        representative K+ channel sequence
  <220>
  <223> eag
  <400> 20
  Tyr Val Thr Ala Leu Tyr Phe Thr Met Thr Cys Met Thr Ser Val Gly
                    5
                                        10
                                                             15
  Phe Gly Asn Val Ala Ala Glu Thr Asp Asn Glu
               20
                                    25
  <210> 21
  <211> 27
```

```
<212> PRT
  <213> Unknown
  <220>
  <223> Description of Unknown Organism: P domain of
        representative K+ channel sequence
  <220>
  <223> ROMK1
  <400> 21
  Met Thr Ser Ala Phe Leu Phe Ser Leu Glu Thr Gln Val Thr Ile Gly
  Tyr Gly Phe Arg Phe Val Thr Glu Gln Cys Ala
               20
  <210> 22
  <211> 27
  <212> PRT
  <213> Unknown
<223> Description of Unknown Organism: P domain of
        representative K+ channel sequence
ر
(220>
(223> IRK1
-F<400> 22
🖺 Phe Thr Ala Ala Phe Leu Phe Ser Ile Glu Thr Gln Thr Thr Ile Gly
Tyr Gly Phe Arg Cys Val Thr Asp Glu Cys Pro
               20
M
IJ
点<210> 23
<211> 27
  <212> PRT
  <213> Unknown
  <220>
  <223> Description of Unknown Organism: P domain of
        representative K+ channel sequence
  <220>
  <223> GIRK1
  <400> 23
  Phe Pro Ser Ala Phe Leu Phe Phe Ile Glu Thr Glu Ala Thr Ile Gly
  Tyr Gly Tyr Arg Tyr Ile Thr Asp Lys Cys Pro
               20
  <210> 24
  <211> 48
  <212> DNA
```

<213> Artificial Sequence

<220>

<400> 24 caccagcagg taggtgaagg tgcacacgat gagagccaac gtgcgcac

48